



GP1624X

Patent

45198.00013.RCE (CIP1)

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re the Application of:

) Group Art Unit: 1624

Applicant: Erion et al.

) Examiner: McKenzie, T.

Serial No.: 09/518,501

)

Filed: March 3, 2000

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For: NOVEL PHOSPHORUS-  
CONTAINING PRODRUGS

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**TRANSMITTAL LETTER**

Commissioner for Patents  
P. O. Box 1450  
Alexandria, VA 22313-1450

Sir:

Enclosed are the following documents:

- Supplemental Information Disclosure Statement;
- Form PTO/SB/08A;
- Check in the amount of \$180; and
- Return Postcard.

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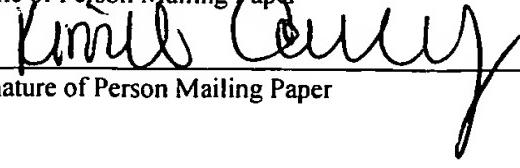
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February 25, 2004  
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The Commissioner is authorized to charge any fee required, or to credit any overpayment, to our Deposit Account No. **50-2613**.

Respectfully submitted,

Dated: 2/25/04

By: Diana L. Bush  
Diana L. Bush, Ph.D., Esq.  
Reg. No. 51,109

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Patent  
45198.00013.RCE (CIP1)

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants: Erion *et al.*

Serial No.: 09/518,501

Filed: March 3, 2000

Title: NOVEL PHOSPHORUS-CONTAINING  
PRODRUGS

Group Art Unit: 1624

Examiner: McKenzie, T.

**SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT**

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P. O. Box 1450  
Alexandria, VA 22313-1450

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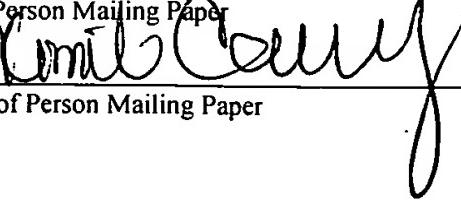
In compliance with Applicants' duty under 37 CFR 1.56, the following information is brought to the attention of the Examiner. The items are listed on the attached Form PTO/SB/08A and copies are enclosed for the convenience of the Examiner. Applicants respectfully request that the documents be made of record in the above-referenced application. Applicants also request that the Examiner return an initialed copy of the attached Form PTO/SB/08A with the next Official Action indicating that the documents have been considered.

The items identified in this Supplemental Information Disclosure Statement may or may not be "material" pursuant to 37 CFR 1.56 and the submission thereof by Applicants shall not be construed as an

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Supplemental Information Disclosure Statement  
Serial No.: 09/518,501  
Page 2 of 2

admission that any such patent, publication or other information referred to therein is material or considered to be material (37 CFR 1.97(h)), or even qualifies as "prior art" under 35 U.S.C. § 102 with respect to this invention unless specifically designated by Applicants as such.

The filing of this Supplemental Information Disclosure Statement shall not be construed to mean that a search has been made or that no other material information, as defined in 37 CFR 1.56, exists.

This Supplemental Information Disclosure Statement is being submitted under 37 CFR 1.97(c)(2). Enclosed is a check including the amount of \$180.00 to cover the fee pursuant to 37 CFR 1.17(p). However, if counsel for Applicants is in error in this regard, the Commissioner is authorized to charge any additional fee, or to credit any overpayment, to Deposit Account No. **50-2613**.

Respectfully submitted,

Dated: 2/25/04

By: Diana L. Bush  
Diana L. Bush, Ph.D., Esq.  
Reg. No. 51,109

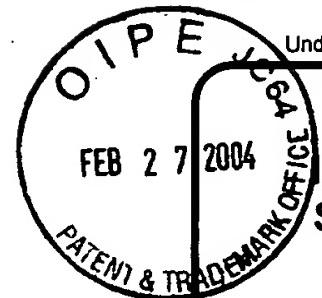
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of

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|------------------------|-------------------------|
| Application Number     | 09/518,501              |
| Filing Date            | March 3, 2000           |
| First Named Inventor   | Erion et al.            |
| Group Art Unit         | 1624                    |
| Examiner Name          | T. McKenzie             |
| Attorney Docket Number | 032465.00013.RCE (CIP1) |

### U.S. PATENT DOCUMENTS

| Examiner Initials* | Cite No. <sup>1</sup> | U.S. Patent Document |                                   | Name of Patentee or Applicant of Cited Document | Date of Publication of Cited Document MM-DD-YYYY | Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear |
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|-------------------|-----------------------|--|----------------|
|                   |                       | Beaucage and Iyer, "The Synthesis of Modified Oligonucleotides by the Phosphoramidite Approach and Their Applications," <u>Tetrahedron</u> , 49(28):6123-6194 (1993).  |                |
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sheet

2 of 3

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| Examiner Name          | T. McKenzie             |
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|-------------------|-----------------------|--|
|                   |                       | Friis and Bundgaard, "Prodrugs of Phosphates and Phosphonates: Novel Lipophilic α-acyloxyalkyl Ester Derivatives of Phosphate- or Phosphonate Containing Drugs Masking the Negative Charges of these Groups," <u>Euro. J. Pharm. Sci.</u> , 4:49-59 (1996).  |
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|                   |                       | Ramachandran et al., "Efficient General Synthesis of 1,2- and 1,3-diols in High Enantiomeric Excess via the Intramolecular Asymmetric Reduction of the Corresponding Ketoalkyl Diisopinocampheylborinate Intermediates," <u>Tetrahedron</u> , 38(5):761-764 (1997).                                |

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|-------------------|-----------------------|---|----------------|
|                   |                       | Starrett, Jr. et al., "Synthesis, Oral Bioavailability Determination, and <i>in Vitro</i> Evaluation of Prodrugs of the Antiviral Agent 9-[2-(Phosphonomethoxy)ethyl]adenine (PMEA)," <u>J. Med. Chem.</u> , 37:1857-1864 (1994).   |                |
|                   |                       | Thomson et al., "Synthesis, Bioactivation and Anti-HIV Activity of the Bis(4-acyloxybenzyl) and Mono(4-acyloxybenzyl) Esters of the 5'-monophosphate of AZT," <u>J. Chem. Soc., Perk. Trans. 1</u> , (11):1239-1245 (1993).   |                |
|                   |                       | Weber and Waxman, "Activation of the Anti-cancer Drug Ifosfamide by Rat Liver Microsomal P450 Enzymes," <u>Biochem. Pharm.</u> , 45(8):1685-1694 (1993).  |                |
|                   |                       | Zon et al., "NMR Spectroscopic Studies of Intermediary Metabolites of Cyclophosphamide. A Comprehensive Kinetic Analysis of the Interconversion of <i>cis</i> - and <i>trans</i> -4-Hydroxycyclophosphamide with Aldophosphamide and Concomitant Partitioning of Aldophosphamide Between Irreversible Fragmentation and Reversible Conjugation Pathways," <u>J. Med. Chem.</u> 27:466-485 (1984). |                |

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